

Type	L#	Hits	Search Text	DBs	Time Stamp	Com ments	Error r Defi nitio n	Error r Defi nitio n
1	BRS	L1	2869 (insulin-like adj growth adj factor-1) or IGF-I	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 09:51		0	
2	BRS	L2	459 solubilizing adj compound	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 09:52		0	
3	BRS	L3	53675 arginine or (guanidine adj hydrochloride) or N-acetyl-arginine or guanidinium	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 09:53		0	
4	BRS	L4	1499 3 same solubiliz\$3	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 09:53		0	
5	BRS	L5	14 1 same 4	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 10:30		0	
6	BRS	L6	2 5410026.pn.	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 10:11		0	
7	BRS	L7	684 "12" adj mg/ml	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 10:37		0	

Type	L #	Hits	Search Text	DBs	Time Stamp	Comments	Error Count	Err Deflators
8	BRS	L8	1963 "200" adj mg/ml	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 10:38		0	
9	BRS	L9	1 5 same (7 or 8)	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 10:38		0	
10	BRS	L10	14 shirley adj bret.in.	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 10:38		0	
11	BRS	L11	1 bajwa adj kamaljit.in.	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 10:39		0	
12	BRS	L12	2 (10 or 11) and 5	USPAT; US-PGPUB; EPO; JPO; DERWENT	2003/08/12 10:39		0	

FILE 'MEDLINE' ENTERED AT 10:43:56 12 AUG 2003

FILE 'CAPLUS' ENTERED AT 10:43:56 ON 12 AUG 2003
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FILE 'SCISEARCH' ENTERED AT 10:43:56 ON 12 AUG 2003
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FILE 'AGRICOLA' ENTERED AT 10:43:56 ON 12 AUG 2003

=> s (insulin-like growth factor-1) or IGF-1
L1 29701 (INSULIN-LIKE GROWTH FACTOR-1) OR IGF-1

=> s arginine or guanidine or guanidium
L2 406219 ARGININE OR GUANIDINE OR GUANIDIUM

=> s 12 (p) solubiliz?
L3 2954 L2 (P) SOLUBILIZ?

=> s 11 (p) 13
L4 5 L1 (P) L3

=> duplicate remove 14
DUPLICATE PREFERENCE IS 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH'
KEEP DUPLICATES FROM MORE THAN ONE FILE? Y/(N):n
PROCESSING COMPLETED FOR L4
L5 2 DUPLICATE REMOVE L4 (3 DUPLICATES REMOVED)

=> d 15 1-2 ibib abs

L5 ANSWER 1 OF 2 CAPLUS COPYRIGHT 2003 ACS on STN
ACCESSION NUMBER: 1999:325815 CAPLUS
DOCUMENT NUMBER: 130:343031
TITLE: Compositions providing for increased IGF-I solubility
INVENTOR(S): Shirley, Bret A.; Bajwa, Kamaljit
PATENT ASSIGNEE(S): Chiron Corporation, USA
SOURCE: PCT Int. Appl., 32 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 3
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9924063	A1	19990520	WO 1998-US23673	19981106
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, CZ, DE, DE, DK, DK, EE, EE, ES, FI, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
AU 9915193	A1	19990531	AU 1999-15193	19981106
EP 1028748	A1	20000823	EP 1998-959383	19981106
EP 1028748	B1	20030502		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
JP 2001522814	T2	20011120	JP 2000-520151	19981106
AT 238807	E	20030515	AT 1998-959383	19981106

PRIORITY APPLN. INFO.: US 1997-64891P P 19971107
WO 1998-US23673 W 19981106

AB IGF-I compns. include a solubilizing compd. comprising a guanidinium group
that provides for IGF-I compns. in which IGF-I is highly sol. at pHs of
about 5.5 or greater and at refrigerated temps. IGF-I was formulated with
arginine for injection.

REFERENCE COUNT:

4

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE SAME FORMAT

L5 ANSWER 2 OF 2 MEDLINE on STN DUPLICATE 1
ACCESSION NUMBER: 92316967 MEDLINE
DOCUMENT NUMBER: 92316967 PubMed ID: 1618780
TITLE: Enhanced insulin-induced mitogenesis and mitogen-activated protein kinase activities in mutant insulin receptors with substitution of two COOH-terminal tyrosine autophosphorylation sites by phenylalanine.
AUTHOR: Ando A; Momomura K; Tobe K; Yamamoto-Honda R; Sakura H; Tamori Y; Kaburagi Y; Koshio O; Akanuma Y; Yazaki Y; +
CORPORATE SOURCE: Third Department of Internal Medicine, Faculty of Medicine, University of Tokyo, Japan.
SOURCE: JOURNAL OF BIOLOGICAL CHEMISTRY, (1992 Jun 25) 267 (18) 12788-96.
PUB. COUNTRY: Journal code: 2985121R. ISSN: 0021-9258.
DOCUMENT TYPE: United States
LANGUAGE: Journal; Article; (JOURNAL ARTICLE)
FILE SEGMENT: English
ENTRY MONTH: Priority Journals
199208
ENTRY DATE: Entered STN: 19920815
Last Updated on STN: 20000303
Entered Medline: 19920806

AB We have studied the function of a mutant human insulin receptor in which two COOH-terminal autophosphorylation sites (Tyr-1316 and -1322) were replaced by phenylalanine (F/Y COOH-terminal 2 tyrosines (CT2)). In addition, we have also constructed a mutant receptor in which Lys-1018 in the ATP-binding site was changed to ***arginine*** (R/K 1018). Both the wild type insulin receptor (HIR) and the mutant receptors were expressed in Chinese hamster ovary (CHO) cells by stable transfection. Autophosphorylation of ***solubilized*** and partially purified F/Y CT2 was decreased by approximately 30% compared with the HIR. Tyrosine kinase activities of F/Y CT2 and HIR toward exogenous substrates were almost equal. When CHO cells transfected with F/Y CT2 (CHO-F/Y CT2) were stimulated with insulin, autophosphorylation of the beta-subunit of the insulin receptor and the phosphorylation of an endogenous substrate (pp185) in the intact cell were normal compared with cells expressing HIR (CHO-HIR). CHO-F/Y CT2 exhibited the same insulin sensitivity as CHO-HIR with respect to 2-deoxyglucose uptake. However, the dose-response curve of insulin-stimulated thymidine incorporation in CHO-F/Y CT2 was shifted to the left (approximately 5-7-fold) compared with that in CHO-HIR. There was no significant difference in ***insulin*** - ***like*** ***growth*** ***factor*** ***1*** -stimulated thymidine incorporation between CHO-F/Y CT2 and CHO-HIR. Furthermore, the dose-response curve of insulin-stimulated kinase activity toward myelin basic protein in CHO-F/Y CT2 was also shifted to the left (approximately 5-fold) compared with that in CHO-HIR. Kinase assays in myelin basic protein-containing gels revealed that both species of MAP kinases (M(r) 44,000, 42,000) were more sensitive to activation by insulin in CHO-F/Y CT2 than in CHO-HIR. This observation was confirmed in immune complex kinase assays toward microtubule-associated protein 2 (MAP2) using specific antibodies against mitogen-activated protein (MAP) kinase. R/K 1018 mutant insulin receptors showed an absence of insulin-stimulated kinase activity and CHO cells transfected with R/K 1018 (CHO-R/K 1018) failed to enhance 2-deoxyglucose uptake or thymidine incorporation in response to insulin. In addition, R/K 1018 kinase-defective insulin receptors were unable to mediate insulin-stimulated MAP kinase activation. These data suggest that: 1) tyrosine kinase activity of the insulin receptor is required for activation of insulin-stimulated MAP kinases and 2) phosphorylation of COOH-terminal tyrosine residues may play an inhibitory role in mitogenic signaling through regulation of MAP kinases.

> d his

(FILE 'HOME' ENTERED AT 10:43:34 ON 12 AUG 2003)

FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH, AGRICOLA' ENTERED AT
10:43:56 ON 12 AUG 2003

L1 29701 S (INSULIN-LIKE GROWTH FACTOR-1) OR IGF-1
L2 406219 S ARGinine OR GUANIDINE OR GUANIDIUM
L3 2954 S L2 (P) SOLUBILIZ?
L4 5 S L1 (P) L3
L5 2 DUPLICATE REMOVE L4 (3 DUPLICATES REMOVED)

=> s (12 mg/ml) or (200 mg/ml)
 'ML' IS NOT A VALID FIELD CODE
 L6 0 (12 MG/ML) OR (200 MG/ML)

=> s shirley bret/au
 L7 6 SHIRLEY BRET/AU

=> s bajwa kamaljit/au
 L8 2 BAJWA KAMALJIT/AU

=> s (17 or 18) and 11
 L9 3 (L7 OR L8) AND L1

=> duplicate remove 19
 DUPLICATE PREFERENCE IS 'CAPLUS, BIOSIS'
 KEEP DUPLICATES FROM MORE THAN ONE FILE? Y/(N):n
 PROCESSING COMPLETED FOR L9
 L10 3 DUPLICATE REMOVE L9 (0 DUPLICATES REMOVED)

=> d 110 1-3 ibib abs

L10 ANSWER 1 OF 3 BIOSIS COPYRIGHT 2003 BIOLOGICAL ABSTRACTS INC. on STN
 ACCESSION NUMBER: 2003:313118 BIOSIS
 DOCUMENT NUMBER: PREV200300313118
 TITLE: Method for producing sustained-release formulations.
 AUTHOR(S): ***Shirley, Bret*** ; Hora, Maninder; O'Hagan, Derek;
 Singh, Manmohan
 ASSIGNEE: Chiron Corporation
 PATENT INFORMATION: US 6573238 June 03, 2003
 SOURCE: Official Gazette of the United States Patent and Trademark
 Office Patents, (June 3 2003) Vol. 1271, No. 1, pp. No
 Pagination. <http://www.uspto.gov/web/menu/patdata.html>.
 e-file.
 ISSN: 0098-1133.

DOCUMENT TYPE: Patent
 LANGUAGE: English
 AB Methods for preparing biodegradable microparticles are provided. Also
 provided are microparticles prepared by the method which include
 IGF - ***1*** entrapped therein. The microparticles allow for
 controlled release of ***IGF*** - ***1*** and other polypeptides
 over prolonged periods of time.

L10 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2003 ACS on STN
 ACCESSION NUMBER: 1999:325815 CAPLUS
 DOCUMENT NUMBER: 130:343031
 TITLE: Compositions providing for increased IGF-I solubility
 INVENTOR(S): Shirley, Bret A.; ***Bajwa, Kamaljit***
 PATENT ASSIGNEE(S): Chiron Corporation, USA
 SOURCE: PCT Int. Appl., 32 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 3
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9924063	A1	19990520	WO 1998-US23673	19981106
W:	AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, CZ, DE, DE, DK, EE, EE, ES, FI, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
AU 9915193	A1	19990531	AU 1999-15193	19981106
EP 1028748	A1	20000823	EP 1998-959383	19981106
EP 1028748	B1	20030502		
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI			
JP 2001522814	T2	20011120	JP 2000-520151	19981106
AT 238807	E	20030515	AT 1998-959383	19981106

PRIORITY APPLN. INFO.:

US 1997-64891P P 19971107

WO 1998-US23673 W 19981106

AB IGF-I compns. include a solubilizing compd. comprising a guanidinium group that provides for IGF-I compns. in which IGF-I is highly sol. at pHs of about 5.5 or greater and at refrigerated temps. IGF-I was formulated with arginine for injection.

REFERENCE COUNT:

4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L10 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER: 1999:325813 CAPLUS

DOCUMENT NUMBER: 130:343029

TITLE: Method for producing ***IGF*** - ***1***
sustained-release formulations

INVENTOR(S): ***Shirley, Bret*** ; Hora, Maninder; O'Hagan,
Derek; Singh, Manmohan

PATENT ASSIGNEE(S): Chiron Corporation, USA

SOURCE: PCT Int. Appl., 60 pp.

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9924061	A1	19990520	WO 1998-US23627	19981106
W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
AU 9913841	A1	19990531	AU 1999-13841	19981106
EP 1028746	A1	20000823	EP 1998-957624	19981106
EP 1028746	B1	20030226		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI				
JP 2001522812	T2	20011120	JP 2000-520149	19981106
US 2002013273	A1	20020131	US 1998-187780	19981106
US 6573238	B2	20030603		
AT 233097	E	20030315	AT 1998-957624	19981106
			US 1997-64891P	P 19971107
			US 1998-96066P	P 19980811
			WO 1998-US23627	W 19981106

PRIORITY APPLN. INFO.:

US 1997-64891P P 19971107
US 1998-96066P P 19980811

WO 1998-US23627 W 19981106

AB Methods for prep. biodegradable poly(D,L-lactide-co-glycolide) microparticles are provided. Also provided are microparticles prep'd. by the method which include ***IGF*** - ***1*** entrapped therein. The microparticles allow for controlled release of ***IGF*** - ***1*** and other polypeptides over prolonged periods of time.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> d his

(FILE 'HOME' ENTERED AT 10:43:34 ON 12 AUG 2003)

FILE 'MEDLINE, CAPLUS, BIOSIS, EMBASE, SCISEARCH, AGRICOLA' ENTERED AT 10:43:56 ON 12 AUG 2003

L1 29701 S (INSULIN-LIKE GROWTH FACTOR-1) OR IGF-1
L2 406219 S ARGININE OR GUANIDINE OR GUANIDIUM
L3 2954 S L2 (P) SOLUBILIZ?
L4 5 S L1 (P) L3
L5 2 DUPLICATE REMOVE L4 (3 DUPLICATES REMOVED)
L6 0 S (12 MG/ML) OR (200 MG/ML)
L7 6 S SHIRLEY BRET/AU
L8 2 S BAJWA KAMALJIT/AU
L9 3 S (L7 OR L8) AND L1
L10 3 DUPLICATE REMOVE L9 (0 DUPLICATES REMOVED)

=> log y

COST IN U.S. DOLLARS

FULL ESTIMATED COST

SINCE FILE ENTRY	TOTAL SESSION
41.35	41.56

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

CA SUBSCRIBER PRICE

SINCE FILE

ENTRY

-1.95

TOT.

SESSION

-1.95

STN INTERNATIONAL LOGOFF AT 10:50:32 ON 12 AUG 2003